

12-24-03

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant: Becker *et al.*  
Serial No.: 10/600,009  
Filed: June 18, 2003  
Customer No.: 24961  
Confirmation No.: Unassigned  
For: **GENES AND POLYMORPHISMS ON  
CHROMOSOME 10 ASSOCIATED WITH  
ALZHEIMER'S DISEASE AND OTHER  
NEURODEGENERATIVE DISEASES**  
Art Unit: Unassigned  
Examiner: Unassigned

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**EV 417728482 US**

Date of Deposit December 22, 2003

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12/22/2003  
Date

Michael Lough

TRANSMITTAL LETTER

Commissioner for Patents  
U.S. Patent and Trademark Office  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Sir:

Transmitted herewith are a Supplemental Information Disclosure Statement, and Form PTO-1449 (14 pages) in connection with the above-identified application. Because this Information Disclosure Statement is filed prior to receipt of a first Office Action on the merits in the above-referenced application, no fee is due. However, should it be determined that a fee for filing these papers is required, the Commissioner is authorized to charge Deposit Account Number 50-1213, as stated below:

(X) The Commissioner is hereby authorized to charge the fee for the Information Disclosure Statement and any other fees that may be due under 37 C.F.R. §§1.16-1.17 in connection with this paper or with this application during its entire pendency to Deposit Account No. 50-1213. A duplicate of this sheet is enclosed.

Respectfully submitted,  
HELLER EHRMAN WHITE & McAULIFFE LLP

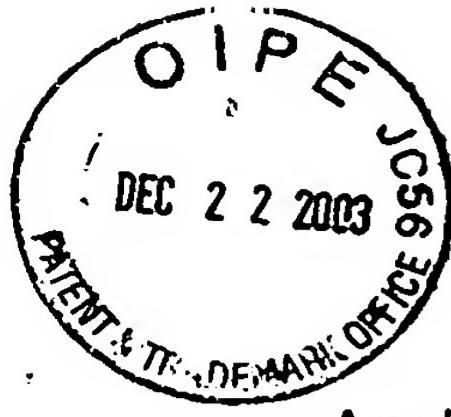
By:

Stephanie L. Seidman  
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Attorney Docket No. 37481-3308B

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### SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT IN ACCORDANCE WITH 37 C.F.R. §§ 1.97-1.98

Commissioner for Patents  
U.S. Patent and Trademark Office  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Sir:

This Supplemental Information Disclosure Statement is filed prior to receipt of a First Office Action on the merits of the above-captioned application. Thus, a fee for filing this statement should not be due. If, however, it is determined that a fee is due, any fees that may be due in connection with filing this paper may be charged to Deposit Account No. 50-1213.

In accordance with the duty of disclosure imposed by 37 C.F.R. §1.56 to inform the Patent Office of all references known by Applicant or Applicant's representative that may be material to the examination of the subject application, Applicant's representative hereby provides this Information Disclosure Statement that is prepared in accordance with 37 C.F.R. §§1.97-1.98. Form PTO-1449 (14 pages) is provided herewith in connection with the above-captioned application. In accordance with 37 C.F.R. §1.98(d), copies of the references marked with an asterisk on the Form PTO-1449 are not provided herewith as they

**U.S.S.N. 10/600,009**

**BECKER *et al.***

**Supplemental Information Disclosure Statement**

have been previously provided in connection with U.S. Serial No. 10/282,174 patent application which is relied upon for an earlier filing date in accordance with 35 U.S.C. §120.

The documents listed on Form PTO-1449 are all in the English language. Hence, in accordance with the requirements of 37 C.F.R. §1.98, as amended effective March 16, 1992, no further explanation of the listed items is necessary.

Applicant also makes known to the Examiner the following U.S. and International applications, which are commonly owned and/or have one or more inventors in common:

<b><u>U.S.S.N.</u></b>	<b><u>Filing Date</u></b>	<b><u>Docket No.</u></b>
09/814,179	03/22/01	37481-3301
10/342,844	01/13/03	37481-3307
10/282,174	10/25/02	37481-3308
10/281,456	10/25/02	37481-3312
60/448,172	02/14/03	37481-P3322
60/451,958	03/04/03	37481-P3322B
10/292,081	11/08/02	37481-3323
10/608,397	06/26/03	37481-3323B
60/470,884	05/14/03	37481-P3327
60/489,186	07/21/03	37481-P3329
<b><u>Int'l App. No.</u></b>	<b><u>Filing Date</u></b>	<b><u>Docket No.</u></b>
PCT/US03/26409	08/20/03	37481-3307CPC
PCT/US02/34679	10/25/02	37481-3308PC
PCT/US03/26173	08/20/03	37481-3316PC

Although these documents are made known to the Patent and Trademark Office in compliance with Applicant's duty of disclosure, such disclosure is not to be construed as an admission by Applicant or Applicant's representative that any of the references, singly or in any combination thereof, is effective as prior art against the subject application. In accordance with 37 C.F.R. §1.97(h), the filing of this Supplemental Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 C.F.R. §1.56(b) exists.

**U.S.S.N. 10/600,009**

**BECKER *et al.***

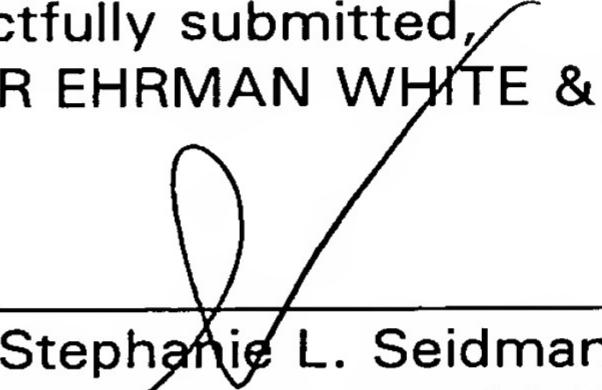
**Supplemental Information Disclosure Statement**

Applicant respectfully requests that the Examiner review the foregoing references and information and that they be made of record in the file history of the above-captioned application.

\* \* \*

Respectfully submitted,  
HELLER EHRMAN WHITE & MCAULIFFE LLP

By:

  
Stephanie L. Seidman  
Registration No. 33,779

Attorney Docket No. 37481-3308B

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FORM PTO-1449	DEC 22 2003 USPTO PATENT & TRADEMARK OFFICE SCS	ATTY. DOCKET NO. 37481-3308B	SERIAL NO. 10/600,009	CONFIRM NO. Unassigned
LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT		APPLICANT Becker <i>et al.</i>	CUSTOMER NO. 24961	
		FILING DATE 06/18/2003	GROUP Unassigned	

\* If an asterisk is placed beside the reference number, a copy is NOT provided because the reference was previously cited by or submitted to the PTO in a prior application that is identified in the statement and relied upon for an earlier filing date under 35 U.S.C. 120. 37 C.F.R. § 1.98(d).

### U.S. PATENT DOCUMENTS

EXAMINER INITIAL	Ref. No.	DOCUMENT NUMBER								DATE	NAME	CLASS	SUB CLASS	FILING DATE
	A*	6	1	5	3	3	8	6	11/28/00	Lalouel <i>et al.</i>	435	6	06/29/98	
	B*	6	3	4	2	3	5	1	01/29/02	Chen <i>et al.</i>	435	6	03/16/99	
	C*	6	3	4	6	3	8	2	02/12/02	Summar <i>et al.</i>	435	6	06/01/99	
	D*	6	3	4	6	3	8	5	02/12/02	Eguchi <i>et al.</i>	435	6	12/06/98	
	E*	6	4	3	2	6	4	8	08/13/02	Blumefeld <i>et al.</i>	435	6	08/16/00	
	F*	6	4	4	4	4	2	7	9/03/02	Ludwig <i>et al.</i>	435	6	07/11/00	

### FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL	Ref. No.	DOCUMENT NUMBER								DATE	COUNTRY	Translation Yes      No
	G*	0	2	0	2	8	2	0	01/10/02	PCT		
	H*	0	3	0	5	1	1	74	06/26/03	PCT A2		
	I*	0	3	0	5	4	1	43	07/03/03	PCT A2		
	J*	9	8	4	7	9	1	7	09/18/98	PCT A2/A3		

### OTHER ART (Including Author, Title, Date, Pertinent Pages, Etc.)

K*	Abraham <i>et al.</i> , "Insulin degrading enzyme is not associated with late-onset Alzheimer's disease," <i>Am. J. Hum. Genet.</i> <u>59</u> : S562, Program No. 2236 (2001), [also available online at <a href="http://www.faseb.org/cgi-bin/ashg01">www.faseb.org/cgi-bin/ashg01</a> , accessed on 07/01/03]
L*	Ait-Ghezala <i>et al.</i> , "Confirmation of association between D10S583 and Alzheimer's disease in a case-control sample," <i>Neuroscience Letters</i> <u>325</u> : 87-90 (2002)

EXAMINER	DATE CONSIDERED
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EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

Title: Genes and Polymorphisms on Chromosome 10 Associated with Alzheimer's Disease and Other Neurodegenerative Diseases

FORM PTO-1449  LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT	DEC 22 2003 PATENT & TRADEMARK OFFICE O I P E C 56	ATTY. DOCKET NO. 37481-3308B	SERIAL NO. 10/600,009	CONFIRM NO. Unassigned
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### OTHER ART (Including Author, Title, Date, Pertinent Pages, Etc.)

M*	American College of Medical Genetics/American Society of Human Genetics (ACMG/ASHG) Working Group on ApoE and Alzheimer Disease, "Statement on use of apolipoprotein E testing for Alzheimer disease. American College of Medical Genetics/American Society of Human Genetics Working Group on ApoE and Alzheimer disease" <i>JAMA</i> <u>274</u> (20):1627-1629 (1995)
N*	American Type Culture Collection, ATCC Number: CRL-9096
O*	Bales <i>et al.</i> , "Lack of apolipoprotein E dramatically reduces amyloid $\beta$ -peptide deposition", <i>Nature Genetics</i> <u>17</u> :263-264 (1997)
P*	Bergen <i>et al.</i> , "The Role of Heredity in Late-Onset Alzheimer Disease and Vascular Dementia", <i>Arch Gen Psychiatry</i> <u>54</u> :264-270 (1997)
Q*	Bertram <i>et al.</i> , "Further Assessment of Novel Alzheimer's Disease Loci on Chromosomes 10 and 9," <i>Neurobiology of Aging</i> <u>23</u> : S324, abstract no. 1206 (2002)
R*	Bertram <i>et al.</i> , "Genetic Analyses of Novel Alzheimer's Disease Loci on Chromosomes 10 and 9," <i>Society for Neuroscience Abstracts</i> Vol. 28, Program No. 420.5 (2002)
S*	Bertram, Lars, "Candidate Genes Showing No Association with Alzheimer's Disease - Results of the NIMH Genetics Initiative," Slides (25) from the presentation at <i>The 2000 World Alzheimer's Congress</i> Washington D.C., July 2000
T*	Bertram, Lars, "Conclusion: Candidate Gene Assessment," Slides (2) from the presentation at <i>The 2000 World Alzheimer's Congress</i> Washington, D.C., July 2000
U*	Bertram, Lars, "Further Characterization of the Alzheimer's Disease Locus on Chromosome 10," Slides (15) from the platform presentation at <i>The 2001 Society for Neuroscience Annual Meeting</i> November 11, 2001
V*	Bertram, Lars, "Genetic Analyses of Novel Alzheimer's Disease Loci on Chromosome 10 and 9," Slides (19) from the platform presentation at <i>The 2002 Society for Neuroscience Annual Meeting</i> November 5, 2002
W*	Bertram, Lars <i>et al.</i> , "Further Assessment of a Novel Alzheimer's Disease Loci on Chromosome 10 and 9," Poster presented at <i>The 2002 International Conference on Alzheimer's Disease and Related Disorders</i> Stockholm, July 2002
X*	Bertram <i>et al.</i> , "Further Characterization of the Alzheimer's Disease Locus on Chromosome 10," <i>Soc. Neurosci. Abstr.</i> Volume 27, Program No. 127.3 (2001)

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#### OTHER ART (Including Author, Title, Date, Pertinent Pages, Etc.)

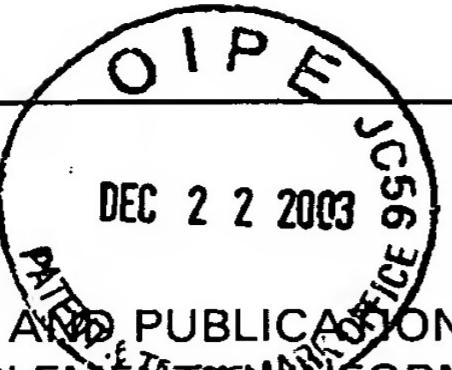
Y*	Bishop, <i>et al.</i> , "The Power of Identity-by-State Methods for Linkage Analysis", <i>Am. J. Hum. Genet.</i> <u>46</u> :254-265 (1990)
Z*	Blacker, Deborah, "Association Strategies in Alzheimer's Disease: Findings from the NIMH Genetics Initiative," Slides (17) from the presentation at <i>The 2000 World Alzheimer's Congress</i> Washington, D.C., July 2000
AA*	Blacker <i>et al.</i> , "ApoE-4 and age at onset of Alzheimer's disease: The NIMH Genetics Initiative", <i>Neurology</i> <u>48</u> :139-147 (1997)
AB*	Blacker <i>et al.</i> , "Results of a high resolution genome screen in 443 Alzheimer's disease families: the NIMH Genetics Initiative," <i>Am. J. Hum. Genet.</i> <u>69</u> : S498, Program No. 1851 (2001) [also available online at <a href="http://www.faseb.org/cgi-bin/ashg01">www.faseb.org/cgi-bin/ashg01</a> , accessed on 07/01/03]
AC*	Blacker <i>et al.</i> , "The Genetics of Alzheimer Disease", <i>Arch Neurol</i> <u>55</u> :294-296 (1998)
AD*	Boehnke <i>et al.</i> , "Genetic Association Mapping Based on Discordant Sib Pairs: The Discordant-Alleles Test", <i>Am. J. Hum. Genet.</i> <u>62</u> :950-961 (1998)
AE*	Breitner <i>et al.</i> , "Familial aggregation in Alzheimer's disease: Comparison of risk among relatives of early- and late-onset cases, and among male and female relatives in successive generations", <i>Neurology</i> <u>38</u> :207-212 (1988)
AF*	Bullido <i>et al.</i> , "A polymorphism in the regulatory region of APOE associated with risk for Alzheimer's dementia", <i>Nature Genetics</i> <u>18</u> :69-71 (1998)
AG*	Chung <i>et al.</i> , "Lack of Association of Trinucleotide Repeat Polymorphisms in the Very-Low-Density Lipoprotein Receptor Gene with Alzheimer's Disease", <i>Ann. Neurology</i> <u>39</u> (6):800-803 (1996)
AH*	Clark, R.F. <i>et al.</i> , Alzheimer's Disease Collaborative Group, "The structure of the presenilin 1 (S182) gene and identification of six novel mutations in early onset AD families", Alzheimer's Disease Collaborative Group, <i>Nat Genet.</i> <u>11</u> (2):219-222 (1995)
AI*	Claworthly <i>et al.</i> , "Lack of Association of a Polymorphism in the Low-density Lipoprotein Receptor-Related Protein Gene With Alzheimer Disease", <i>Arch Neurol</i> <u>54</u> :1289-1292 (1997)
AJ*	Clerget-Darpoux <i>et al.</i> , "Effects of Misspecifying Genetic Parameters in Lod Score Analysis", <i>Biometrics</i> <u>42</u> :393-399 (1986)
AK*	Cook <i>et al.</i> , "Twins With Alzheimer's Disease", <i>Arch Neurol</i> <u>38</u> :3001-301 (1981)

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AL*	Corder <i>et al.</i> , "Apolipoprotein E, survival in Alzheimer's disease patients, and the competing risks of death and Alzheimer's disease", <i>Neurology</i> <u>45</u> :1323-1328 (1995)
AM*	Creasey <i>et al.</i> , "Monozygotic twins discordant for Alzheimer's disease", <i>Neurology</i> <u>39</u> :1474-1476 (1989)
AN*	Cruts <i>et al.</i> , "Estimation of the genetic contribution of presenilin-1 and -2 mutations in a population-based study of presenile Alzheimer disease", <i>Hum Mol Genetics</i> <u>7</u> (1):43-51 (1998)
AO*	Cruts <i>et al.</i> , "The presenilin genes: a new gene family involved in Alzheimer disease pathology", <i>Hum Mol Genetics</i> <u>5</u> :1449-1455 (1996)
AP*	Du <i>et al.</i> , " $\alpha_2$ -Macroglobulin as a $\beta$ -Amyloid Peptide-Binding Plasma Protein", <i>J. Neurochemistry</i> <u>69</u> (1):299-305 (1997)
AQ*	Du <i>et al.</i> , " $\alpha_2$ -Macroglobulin Attenuates $\beta$ -Amyloid Peptide 1-40 Fibril Formation and Associated Neurotoxicity of Cultured Fetal Rat Cortical Neurons", <i>J. Neurochemistry</i> <u>70</u> (3):1182-1188 (1998)
AR*	Elston <i>et al.</i> , "Two-Stage Global Search Designs for Linkage Analysis Using Pairs of Affected Relatives", <i>Genetic Epidemiology</i> <u>13</u> :535-558 (1996)
AS*	Elston, R.C., "Designs for the Global Search of the Human Genome by Linkage Analysis", Proceedings of the XVIth International Biometric Conference, Hamilton, New Zealand, pp. 39-51 (1992)
AT*	Ertekin-Taner <i>et al.</i> , "Analysis of Urokinase-Type Plasminogen Activator as a Late-Onset Alzheimer's Disease Candidate Gene Linked To Chromosome 10," <i>51st Annual Meeting of the American Society of Human Genetics</i> San Diego, California, USA (October 2001)
AU*	Ertekin-Taner <i>et al.</i> , "Significant Association of Haplotype Blocks in the Urokinase Type Plasminogen Activator Gene on Chromosome 10 with $\beta$ 42 and Late Onset Alzheimer's Disease," from the 52nd Annual Meeting of the American Society of Human Genetics Baltimore, Maryland USA October 15-19, 2002. Published in <i>American Journal of Human Genetics</i> <u>71</u> (4 Supplement): 458 October, 2002
AV*	Ertekin-Taner <i>et al.</i> , "Urokinase-type plasminogen activator haplotypes are significantly associated with $\beta$ 42 and late onset Alzheimer's disease," <i>Presentations at Neuroscience Symposium "The Latest Links: Genes and Alzheimer's Disease." National Institute of Aging Neuroscience and Neuropsychology of Aging</i> in connection with <i>The 2001 Society for Neuroscience Annual Meeting</i> November 10, 2001 San Diego, CA

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#### OTHER ART (Including Author, Title, Date, Pertinent Pages, Etc.)

AW *	Ertekin-Taner <i>et al.</i> , "Variance components linkage analysis of plasma amyloid $\beta$ protein in typical late-onset Alzheimer's disease (LOAD) pedigrees," <i>Am. J. Hum. Genet.</i> <u>69</u> : S531, Program No. 2048 (2000) [also available online at <a href="http://www.faseb.org/cgi-bin/ashg01">www.faseb.org/cgi-bin/ashg01</a> , accessed on 07/01/03]
AX *	Estus <i>et al.</i> , "Urokinase-type Plasminogen Activator, Amyloid-beta, and Alzheimers Disease," <i>Presentations at Neuroscience Symposium "The Latest Links: Genes and Alzheimer's Disease."</i> National Institute of Aging Neuroscience and Neuropsychology of Aging in connection with <i>The 2001 Society for Neuroscience Annual Meeting</i> November 10, 2001 San Diego, CA
AY *	Evans <i>et al.</i> , "Apolipoprotein E $\epsilon$ 4 and Incidence of Alzheimer Disease in a Community Population of Older Persons", <i>JAMA</i> <u>277</u> (10):822-824 (1997)
AZ *	Evans <i>et al.</i> , "Prevalence of Alzheimer's Disease in a Community Population of Older Persons", <i>JAMA</i> <u>262</u> (18):2551-2556 (1989)
BA *	Farrer <i>et al.</i> , "Assessment of Genetic Risk for Alzheimer's Disease Among First-Degree Relatives", <i>Ann. Neurology</i> <u>25</u> (5):485-493 (1989)
BB *	Farrer <i>et al.</i> , "Transmission and age-at-onset patterns in familial Alzheimer's disease: Evidence for heterogeneity", <i>Neurology</i> <u>40</u> :395-403 (1990)
BC *	Feingold <i>et al.</i> , "Strategies for Mapping Heterogeneous Recessive Traits by Allele-Sharing Methods", <i>Am. J. Genet.</i> <u>60</u> :965-978 (1997)
BD *	Feuk <i>et al.</i> , "Multiple Genes Maybe Involved in Explaining the Linkage Peak on Chromosome 10: Evidence From SNP-Based Association Studies," <i>Neurobiology of Aging</i> <u>23</u> : S424-S425 (2002)
BE *	Games <i>et al.</i> , "Alzheimer-type neuropathology in transgenic mice over expressing V717F $\beta$ -amyloid precursor protein", <i>Nature</i> <u>373</u> :523-527 (1995)
BF *	Go <i>et al.</i> , "Development and Validation of a Structured Telephone Interview for Dementia Assessment (STIDA): The NIMH Genetics Initiative", <i>J. Geriatr. Psychiatry Neurol</i> <u>10</u> :161-167 (1997)
BG *	Goldin <i>et al.</i> , "Improvement of the Power to Detect Complex Disease Genes by Regional Inference Procedures", <i>Genetic Epidemiology</i> <u>14</u> :785-789 (1997)
BH *	Gómez-Isla <i>et al.</i> , "A Novel Presenilin-1 Mutation: Increased $\beta$ -Amyloid and Neurofibrillary Changes", <i>Ann. Neurol</i> <u>41</u> :809-813 (1997)

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BI *	Goudsmit <i>et al.</i> , "Familial Alzheimer's Disease in Two Kindreds of the Same Geographic and Ethnic Origin", <i>J Neurology Sciences</i> <u>49</u> :79-89 (1981)
BJ *	Greenberg <i>et al.</i> , "Affecteds-Only Linkage Methods Are Not a Panacea", <i>Am. J. Hum. Genet.</i> <u>58</u> :892-895 (1996)
BK *	Growdon <i>et al.</i> , NIH Grant 5P50AG005134-20 "Massachusetts Alzheimer's Disease Research Center," funding period 04/01/1999-03/21/2004
BL *	Hardy, J., "Amyloid, the presenilins and Alzheimer's disease", <i>20</i> :154-159 (1997)
BM *	Haseman <i>et al.</i> , "The Investigation of Linkage Between a Quantitative Trait and a Marker Locus", <i>Behavior Genetics</i> <u>2</u> (1):3-19(1972)
BN *	Hauser <i>et al.</i> , "Affected-Sib-Pair Interval Mapping and Exclusion for Complex Genetic Traits: Sampling Considerations", <i>Genetic Epidemiology</i> <u>13</u> :117-137 (1996)
BO *	Hawley <i>et al.</i> , "A computer program implementing the EM algorithm for haplotype frequency estimation", <i>American Journal of Physical Anthropology Conference/Meeting: 63rd Annual Meeting of the American Association of Physical Anthropologists, Denver, Colorado, USA, March 29-April 2, 1994, AAPA Abstracts, Suppl 18, p. 104 (1994)</i>
BP *	Heath <i>et al.</i> , "Markov Chain Monte Carlo Segregation and Linkage Analysis for Oligogenic Models", <i>Am. J. Hum. Genet.</i> <u>61</u> :748-760 (1997)
BQ *	Helbecque <i>et al.</i> , "The Role of a Triplet Repeat Sequence of the Very Low Density Lipoprotein Receptor Gene in Plasma Lipid and Lipoprotein Level Variability in Humans", <i>Arteriosclerosis</i> <u>17</u> (11):2759-2764 (1997)
BR *	Hendriks <i>et al.</i> , "Presenile dementia and cerebral haemorrhage linked to a mutation at codon 692 of the $\beta$ -amyloid precursor protein gene", <i>Nature Genetics</i> <u>1</u> :218-221 (1992)
BS *	Hodge <i>et al.</i> , "Magnitude of Type I Error When Single-Locus Linkage Analysis Is Maximized over Models: A Simulation Study", <i>Am. J. Hum. Genet.</i> <u>60</u> :217-227 (1997)
BT *	Hollenbach <i>et al.</i> , "Confirmation of an association between a polymorphism in exon 3 of the low-density lipoprotein receptor-related protein gene and Alzheimer's disease", <i>Neurology</i> <u>50</u> (6):1905-1907 (1998)
BU *	Holmans <i>et al.</i> , "Asymptotic Properties of Affected-Sib-Pair Linkage Analysis", <i>Am. J. Hum. Genet.</i> <u>52</u> :362-374 (1993)
BV *	Hsiao <i>et al.</i> , "Correlative Memory Deficits, A $\beta$ Elevation, and Amyloid Plaques in Transgenic Mice", <i>Science</i> <u>274</u> :99-102 (1996)

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Title : Genes and Polymorphisms on Chromosome 10 Associated with Alzheimer's Disease and Other Neurodegenerative Diseases

FORM PTO-1449  LIST OF PATENTS AND PUBLICATIONS FOR APPLICANT'S SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 37481-3308B	SERIAL NO. 10/600,009	CONFIRM NO. Unassigned
	APPLICANT <i>Becker et al.</i>	CUSTOMER NO. 24961	
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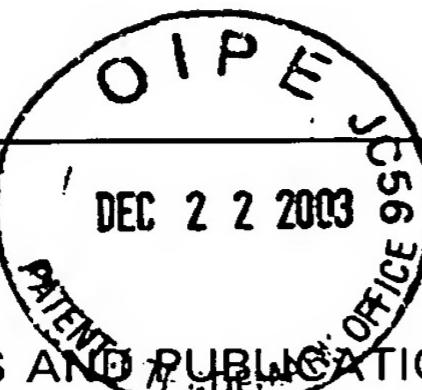
BW*	Huff <i>et al.</i> , "Age at Onset and Rate of Progression of Alzheimer's Disease", <i>JAGS</i> <u>35</u> :27-30 (1987)
BX*	Huff <i>et al.</i> , "Risk of dementia in relatives of patients with Alzheimer's disease", <i>Neurology</i> <u>38</u> :786-790 (1988)
BY*	Hughes <i>et al.</i> , " $\alpha_2$ -macroglobulin associates with $\beta$ -amyloid peptide and prevents fibril formation", <i>Proc. Natl. Acad. Sci. USA</i> <u>95</u> :3275-3280 (1998)
BZ*	Hutton <i>et al.</i> , "Complete analysis of the presenilin 1 gene in early onset Alzheimer's disease", <i>NeuroReport</i> <u>7</u> :801-805 (1996)
CA*	Hyman <i>et al.</i> , "Apolipoprotein E and Cognitive Change in an Elderly Population", <i>Ann Neurology</i> <u>40</u> (1):55-66 (1996)
CB*	Jarvik <i>et al.</i> , "Organic Brain Syndrome and Aging: A Six-Year Follow-up of Surviving Twins", <i>Arch Gen Psychiatry</i> <u>37</u> :280-286 (1980)
CC*	Katzman <i>et al.</i> , "The Epidemiology of Dementia and Alzheimer Disease", Book: <u>Alzheimer Disease</u> , Raven Press, Ltd., New York, Chapter 8, pp. 105-122 (1994)
CD*	Kaufman <i>et al.</i> , "Evolution of Chromosomal Regions Containing Transfected and Amplified Dihydrofolate Reductase Sequences", <i>Mol Cell Biol</i> <u>3</u> (4):699-711 (1983)
CE*	Khachaturian, Z.S., "Diagnosis of Alzheimer's Disease", <i>Arch Neurol</i> <u>42</u> :1097-1105 (1985)
CF*	Knapp <i>et al.</i> , "Linkage Analysis in Nuclear Families", <i>Hum Hered</i> <u>44</u> :37-43 (1994)
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CH*	Kounnas <i>et al.</i> , "LDL Receptor-Related Protein, a Multifunctional ApoE Receptor, Binds Secreted $\beta$ -Amyloid Precursor Protein and Mediates Its Degradation", <i>Cell</i> <u>82</u> :31-340 (1995)
CI*	Kovacs <i>et al.</i> , "Alzheimer-associated presenilins 1 and 2: Neuronal expression in brain and localization to intracellular membranes in mammalian cells", <i>Nature Medicine</i> <u>2</u> (2):224-229 (1996)
CJ*	Kruglyak <i>et al.</i> , "Parametric and Nonparametric Linkage Analysis: A Unified Multipoint Approach", <i>Am. J. Hum. Genet.</i> <u>58</u> :1347-1363 (1996)

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CL*	Lambert <i>et al.</i> , "Distortion of allelic expression of apolipoprotein E in Alzheimer's disease", <i>Hum Mol Genetics</i> 6(12):2151-2154 (1997)
CM*	Lander <i>et al.</i> , "Construction of multilocus genetic linkage maps in humans", <i>Proc. Natl. Acad. Sci. USA</i> 84:2363-2367 (1987)
CN*	Lautenschlager <i>et al.</i> , "Risk of dementia among relatives of Alzheimer's disease patients in the MIRAGE study: What is in store for the oldest old?", <i>Neurology</i> 46:641-649 (1996)
CO*	Lendon <i>et al.</i> , "Genetic association studies between dementia of the Alzheimer's type and three receptors for apolipoprotein E in a Caucasian population", <i>Neuroscience Letters</i> 222:187-190 (1997)
CP*	Levy <i>et al.</i> , "Mutation of the Alzheimer's Disease Amyloid Gene in Hereditary Cerebral Hemorrhage, Dutch Type", <i>Science</i> 248:1124-1126 (1990)
CQ*	Levy-Lahad <i>et al.</i> , "Candidate Gene for the Chromosome 1 Familial Alzheimer's Disease Locus", <i>Science</i> 269:973-977 (1995)
CR*	Li <i>et al.</i> , "Genome-wide scan for age at onset of Alzheimer's disease," <i>Am. J. Hum. Genet.</i> 69: S200, Program No. 126 (2001) [also available online at <a href="http://www.faseb.org/cgi-bin/ashg01">www.faseb.org/cgi-bin/ashg01</a> , accessed on 07/01/03]
CS*	Luxenberg <i>et al.</i> , "Cerebral metabolism, anatomy, and cognition in monozygotic twins discordant for dementia of the Alzheimer type", <i>J. Neurology, Neurosurgery, and Psychiatry</i> 50:33-340 (1987)
CT*	Matthijs <i>et al.</i> , "A deletion polymorphism in the human alpha-2-macroglobulin (A2M) gene", <i>Nucleic Acids Res</i> 19(18):5102 (1991)
CU*	Mayeux <i>et al.</i> , "Risk of Dementia in First-Degree Relatives of Patients With Alzheimer's Disease and Related Disorders", <i>Arch Neurol</i> 48:269-273 (1991)
CV*	McGeer <i>et al.</i> , "Anti-inflammatory agents as a therapeutic approach to Alzheimer's disease", <i>Neurology</i> 42:447-449 (1992)
CW*	Meyer <i>et al.</i> , "APOE genotype predicts when-not whether-one is predisposed to develop Alzheimer disease", <i>Nature Genetics</i> 19:321-322 (1998)

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CY*	Montoya <i>et al.</i> , "Bleomycin hydrolase is associated with risk of sporadic Alzheimer's disease", <i>Nature Genetics</i> <u>18</u> :211-212 (1998)
CZ*	Mullan <i>et al.</i> , "A pathogenic mutation for probable Alzheimer's disease in the APP gene at the N-terminus of $\beta$ -amyloid", <i>Nature Genetics</i> <u>1</u> :345-347 (1992)
DA*	Mullin <i>et al.</i> , "Linkage Analysis in a Large Late-Onset Familial Alzheimer's Disease Pedigree," <i>Soc. Neurosci. Abstr.</i> <u>Vol. 27</u> , Program No. 192.4 (2001)
DB*	Myers <i>et al.</i> , "Apolipoprotein E $\epsilon$ 4 association with dementia in a population-based study: The Framingham Study", <i>Neurology</i> <u>46</u> :673-677 (1996)
DC*	Myers <i>et al.</i> , "Progress Toward Identification of the chromosome 10 locus for late onset Alzheimer's disease," <i>Presentations at Neuroscience Symposium "The Latest Links: Genes and Alzheimer's Disease." National Institute of Aging Neuroscience and Neuropsychology of Aging in connection with The 2001 Society for Neuroscience Annual Meeting November 10, 2001 San Diego, CA</i>
DD*	Nagamine <i>et al.</i> , "Structure and Function of the Urokinase-Type Plasminogen Activator Gene", Book: <u>Fibrinolysis in Disease: Molecular and Hemovascular Aspects of Fibrinolysis</u> , CRC Press, Boca Raton, Chapter 2, pp. 10-20 (1996)
DE*	Namekata <i>et al.</i> , "Association of transferrin C2 allele with late-onset Alzheimer's disease", <i>Hum Genet</i> <u>101</u> :126-129 (1997)
DF*	Narita <i>et al.</i> , " $\alpha$ 2-Macroglobulin Complexes with and Mediates the Endocytosis of $\beta$ -Amyloid Peptide via Cell Surface Low-Density Lipoprotein Receptor-Related Protein", <i>J. Neurochem</i> <u>69</u> (5):1904-1911 (1997)
DG*	National Institute on Aging/Alzheimer's Association Working Group, "Apolipoprotein E genotyping in Alzheimer's disease", <i>Lancet</i> <u>347</u> (9008):1091-1095 (1996)
DH*	Nee <i>et al.</i> , "A Family With Histologically Confirmed Alzheimer's Disease", <i>Arch Neurol</i> <u>40</u> :203-208 (1983)
DI*	Nee <i>et al.</i> , "Dementia of the Alzheimer type: Clinical and family study of 22 twin pairs", <i>Neurology</i> <u>37</u> :359-363 (1987)

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DJ*	Nowotny <i>et al.</i> , "Analysis of a strong candidate gene for late onset Alzheimer's disease within the 1-lod interval on chromosome 10," from the 52nd Annual Meeting of the American Society of Human Genetics Baltimore, Maryland USA October 15-19, 2002. Published in <i>American Journal of Human Genetics</i> <u>71</u> (4 Supplement): S450 October, 2002
DK*	Payami <i>et al.</i> , "A Prospective Study of Cognitive Health in the Elderly (Oregon Brain Aging Study): Effects of Family History and Apolipoprotein E Genotype", <i>Am. J. Hum. Genet.</i> <u>60</u> :948-956 (1997)
DL*	Payami <i>et al.</i> , "Evidence for association of HLA-A2 allele with onset age of Alzheimer's disease", <i>Neurology</i> <u>49</u> :512-518 (1997)
DM*	Pericak-Vance <i>et al.</i> , "Modulation of Age at Onset and Risk in Alzheimer Disease (AD)," <i>Presentations at Neuroscience Symposium "The Latest Links: Genes and Alzheimer's Disease."</i> <i>National Institute of Aging Neuroscience and Neuropsychology of Aging</i> in connection with <i>The 2001 Society for Neuroscience Annual Meeting</i> November 10, 2001 San Diego, CA
DN*	Perry <i>et al.</i> , "A strong association of the XmnI promoter polymorphism of the gamma globin gene (HBG2) to Alzheimers disease in the NIMH sibling dataset," from the 52nd Annual Meeting of the American Society of Human Genetics Baltimore, Maryland USA October 15-19, 2002. Published in <i>American Journal of Human Genetics</i> <u>71</u> (4 Supplement): 434 October, 2002
DO*	Poller <i>et al.</i> , "Cloning of the human $\alpha_2$ -macroglobulin gene and detection of mutations in two functional domains: the bait region and the thiolester site", <i>Hum Genet</i> <u>88</u> :383-319 (1992)
DP*	Price <i>et al.</i> , "Alzheimer disease-when and why?", <i>Nature Genetics</i> <u>19</u> :314-316 (1998)
DQ*	Qiu <i>et al.</i> , "Degradation of amyloid $\beta$ -Protein by a Serine Protease- $\alpha_2$ -Macroglobulin Complex", <i>J. Biol Chem</i> <u>271</u> (14):8443-8451 (1996)
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DS*	Risch, N., "Exclusion mapping for complex diseases," <i>Am. J. Hum. Genet.</i> <u>53</u> (Suppl): 185 (1993)
DT*	Rogaev <i>et al.</i> , "Familial Alzheimer's disease in kindreds with missense mutations in a gene on chromosome 1 related to the Alzheimer's disease type 3 gene", <i>Nature</i> <u>376</u> :775-778 (1995)

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	DU*	Rosenbaum <i>et al.</i> , "Temperature-gradient gel electrophoresis: Thermodynamic analysis of nucleic acids and proteins in purified form and in cellular extracts", <i>Biophysical Chem</i> <u>26</u> :235-246 (1987)
	DV*	Rosenberg, M.J., "ClinAccess™: An Integrated Client/Server Approach to Clinical Data Management and Regulatory Approval", SUGI 21: Proceedings of the Twenty-First Annual SAS Users Group International Conference, Chicago, Illinois, March 10-13, 1996 pp. 1190-1198.
	DW*	Roses, A.D. and Pericak-Vance, M.A., "Alzheimer's Disease and Other Dementias," <u>Emery and Rimoin's Principles and Practice of Medical Genetics</u> , Vol. II, 3rd Ed., Churchill Livingstone, pp. 8807-1825 (1997)
	DX*	Rosser <i>et al.</i> , "Incomplete penetrance of familial Alzheimer's disease in a pedigree with a novel presenilin-1 gene mutation," <i>The Lancet</i> <u>347</u> :1560 (1996)
	DY*	Schellenberg <i>et al.</i> , "Genetic Linkage Evidence for a Familial Alzheimer's Disease Locus on Chromosome 14", <i>Science</i> <u>258</u> :668-671 (1992)
	DZ*	Schork <i>et al.</i> , "Two-Trait-Locus Linkage Analysis: A Powerful Strategy for Mapping Complex Genetic Traits", <i>Am. J. Hum. Genet.</i> <u>53</u> :1127-1136 (1993)
	EA*	Scott <i>et al.</i> , "No genetic association between the LRP receptor and sporadic or late-onset familial Alzheimer disease", <i>Neurogenetics</i> <u>1</u> :179-183 (1998)
	EB*	Sherrington <i>et al.</i> , "Cloning of a gene bearing missense mutations in early-onset familial Alzheimer's disease", <i>Nature</i> <u>375</u> :754-760 (1995)
	EC*	Smalley <i>et al.</i> , "A General Statistical Model for Detecting Complex-Trait Loci by Using Affected Relative Pairs in a Genome Search", <i>Am. J. Hum. Genet.</i> <u>58</u> :844-860 (1996)
	ED*	Snow <i>et al.</i> , "A Close Ultrastructural Relationship between Sulfated Proteoglycans and AA Amyloid Fibrils", <i>Lab Invest</i> <u>57</u> (6):687-698 (1987)
	EE*	Sorbi <i>et al.</i> , "Epistatic Effect of APP717 Mutation and Apolipoprotein E Genotype in Familial Alzheimer's Disease", <i>Ann Neurology</i> <u>38</u> (1):124-127 (1995)
	EF*	Spielman <i>et al.</i> , "A Sibship Test for Linkage in the Presence of Association: The Sib Transmission/Disequilibrium Test", <i>Am. J. Hum. Genet.</i> <u>62</u> :450-458 1998)
	EG*	Stewart <i>et al.</i> , "The watermaze", Book: <u>Behavioural Neuroscience, Volume I: A Practical Approach</u> , Oxford University Press, Chapter 9, pp. 107-122, (1993)

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EH *	Strickland <i>et al.</i> , "Sequence Identity between the $\alpha_2$ -Macroglobulin Receptor and Low Density Lipoprotein Receptor-related Protein Suggests That This Molecule Is a Multifunctional Receptor", <i>J. Biol. Chem.</i> <u>265</u> (29):17401-17404 (1990)
EI *	Tanzi <i>et al.</i> , "Amyloid $\beta$ Protein Gene: cDNA, mRNA Distribution, and Genetic Linkage Near the Alzheimer Locus", <i>Science</i> <u>235</u> :880-884 (1987)
EJ *	Tanzi <i>et al.</i> , "The Presenilin Genes and Their Role in Early-Onset Familial Alzheimer's Disease", <i>Alzheimer's Disease Review</i> <u>1</u> :90-98 (1996)
EK *	Tanzi <i>et al.</i> , "The Search for Novel Alzheimer's Disease Genes," <i>Neurobiology of Aging</i> <u>23</u> : S10 (2002)
EL *	Tanzi <i>et al.</i> , "The Search for Novel Alzheimer's Disease Genes," <i>Presentations at Neuroscience Symposium "The Latest Links: Genes and Alzheimer's Disease." National Institute of Aging Neuroscience and Neuropsychology of Aging</i> in connection with <i>The 2001 Society for Neuroscience Annual Meeting</i> November 10, 2001 San Diego, CA
EM *	Tanzi <i>et al.</i> , "The Wilson disease gene is a copper transporting ATPase with homology to the Menkes disease gene", <i>Nature Genetics</i> <u>5</u> :344-350 (1993)
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EO *	Tanzi, Rudolph, "Genes linked to Alzheimer's Disease," Slides (38) from the presentation at the <i>Neuroscience Symposium entitled "The Latest Links: Genes and Alzheimer's Disease. National Institute of Aging, Neuroscience and Neuropsychology of Aging"</i> in connection with <i>The 2001 Society for Neuroscience Annual Meeting</i> November 10, 2001
EP *	Tanzi, Rudolph, "Search for Novel AD Genes: Overview," Slides (31) from the presentation at <i>The 2002 International Conference on Alzheimer's Disease and Related Disorders</i> Stockholm, July 20, 2002
EQ *	Terwilliger <i>et al.</i> , "True and False Positive Peaks in Genomewide Scans: Applications of Length-Biased Sampling to Linkage Mapping", <i>Am. J. Hum. Genet.</i> <u>61</u> :430-438 (1997)
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### OTHER ART (Including Author, Title, Date, Pertinent Pages, Etc.)

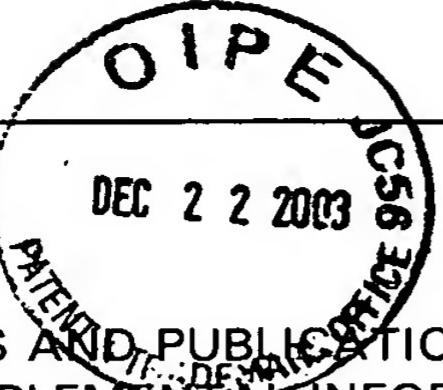
	ET*	Van Broeckhoven, C., "Presenilins and Alzheimer disease", <i>Nature Genetics</i> <u>11</u> :230-232 (1995)
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	EV*	Wavrante-DeVrieze <i>et al.</i> , "Association between the low density lipoprotein receptor-related protein (LRP) and Alzheimer's disease", <i>Neuroscience Letters</i> <u>227</u> :68-70 (1997)
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	EZ*	Wilson <i>et al.</i> , "Apolipoprotein E Alleles, Dyslipidemia, and Coronary Heart Disease", <i>JAMA</i> <u>272</u> (21):1666-1671 (1994)
	FA*	Wilson <i>et al.</i> , "Statistical Validity of the Haseman-Elston Sib-Pair Test in Small Samples", <i>Genetic Epidemiology</i> <u>10</u> :593-598 (1993)
	FB*	Wragg <i>et al.</i> , "Genetic association between intronic polymorphism in presenilin-1 gene and late-onset Alzheimer's disease", <i>The Lancet</i> <u>347</u> :509-512 (1996)
	FC*	Wright, S. Book: <u>Evolution and the Genetics of Populations, Volume 1: Genetic and Biometric Foundations</u> , University of Chicago Press, pp. 411-417 (1968)
	FD*	Younkin <i>et al.</i> , "Identification of a Novel Late Onset Alzheimer's Disease Gene on Chromosome 10," from the 52nd Annual Meeting of the American Society of Human Genetics Baltimore, Maryland USA October 15-19, 2002. Published in <i>American Journal of Human Genetics</i> <u>71</u> (4 Supplement): 215 October, 2002
	FE*	Zhang <i>et al.</i> , "Inhibition of $\alpha_2$ -macroglobulin/proteinase-mediated degradation of amyloid $\beta$ peptide by apolipoprotein E and $\alpha 1$ -antichymotrypsin", <i>Amyloid: Int. J. Exp. Clin. Invest.</i> <u>3</u> :156-161 (1996)

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FORM PTO-1449  DEC 22 2003 APPLICANT Becker et al. FILING DATE 06/18/2003	ATTY. DOCKET NO. 37481-3308B	SERIAL NO. 10/600,009	CONFIRM NO. Unassigned
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	FF*	Zubenko et al., "Monozygotic Twins Concordant for Probable Alzheimer Disease and Increased Platelet Membrane Fluidity", <i>Am. J. Med. Genetics</i> <u>29</u> :431-436 (1988)
	FG *	Zuliani et al., "Tetranucleotide length polymorphism 5' of the $\alpha$ 2-macroglobulin receptor (A2MR)/LDL receptor-related protein (LRP) gene", <i>Human Molecular Genetics</i> <u>3</u> (1):215 (1994)

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